

#3

FIG. 1A

1 aggggagaggc agtgaccatg aaggctgtgc tgccttgccct gttgatggca
 51 ggcttggccc tgcagccagg cactgccctg ctgtgctact cctgcaaagc
 101 ccaggtgagc aacgaggact gcctgcaggt ggagaactgc acccagctgg
 151 gggagcagtg ctggaccgcg cgcattccgcg cagttggcct cctgaccgtc
 201 atcagcaaag gctgcagctt gaactgcgtg gatgactcac aggactacta
 251 cgtgggcaag aagaacatca cgtgctgtga caccgacttg tgcaacgccca
 301 gcggggccca tgccctgcag ccggctgccc ccatccttgc gctgctccct
 351 gcactcggcc tgctgctctg gggaccggcg cagctatagg ctctgggggg
 401 ccccgtgca gcccaactg ggtgtggtgc cccaggcctt tgtgccactc
 451 ctacagaac ctggcccagt gggagcctgt cctggttccct gaggcacatc
 501 ctaacgcaag ttgaccatg tatgtttgca ccccttttcc ccnaaccctg
 551 accttccat gggccttttc caggattccn accnggcaga tcagtttttag
 601 tganacanat ccgcntgcag atggcccctc caacnnttn tgttngtgtt
 651 tccatggccc agcatatttc acccttaacc ctgtgttcag gcactnttcc
 701 cccagggaag cttccctgc ccacccatt tatgaattga gccaggtttg
 751 gtccgtggtg tcccccgcac ccagcagggg acaggcaatc aggaggccc
 801 agtaaaggct gagatgaagt ggactgagta gaactggagg acaagagttg
 851 acgtgagttc ctgggagttt ccagagatgg ggcctggagg cctggaggaa
 901 ggggccaggc ctacatttg tgggntccc gaatggcagc ctgagcacag
 951 cgtaggccct taataaacac ctgttgata agccaaaaaa aaaaaaaa

FIG. 1B

MKAVLLALLMAGLALQPGTALLCYSCKAQVSNECDLQV

ENCTQLGEQCWTARIRAVGLLTVISKGCSLNCVDDS

QDYVVGKKNITCCDIDLNASGAHALQAAAILALLPAL

GLLLWGPGQL

FIG. 2

1 ATGAAGACAGT TTTT TTTATCCTGCTG GGCACCTACTTAGCCCTGCATCCAGGTGCTGCT
 -----+-----+-----+-----+-----+ 60
 TACTTCTGTCAAAAAAAAAATAGGACGACCGGTGGATGAATCGGGACGTAGGTCCACGACGA

 M K T V F F I L L A T Y L A L H P G A A

 CTGCAGTGCATTATCATGCACAGCACAGATGAACAACAGAGACTGCTGTAATGTACAGAAC
 61 -----+-----+-----+-----+-----+ 120
 GACGTCACGATAAGTACGTGCTGCTACTTGTGTCTCTGACAGACTTACATGCTCTTG

 L Q C Y S C T A Q M N N R D C L N V Q N

 TGCAGCCTGGACGACAGTGTGCTTTACATCGCGATCCGGGCCATTGGACTCGTGACA
 121 -----+-----+-----+-----+-----+ 180
 ACGTCGGACCTGGTCTGTCAACGAAATGTAGCGCGTAGGCCCGTAACCTGAGCACTGT

 C S L D Q H S C F T S R I R A I G L V T

 GTTATCAGTAAGGGCTGCAGCTCACAGTGTGAGGATGACTCGGAGAACTACTATTGGGGC
 181 -----+-----+-----+-----+-----+ 240
 CAATAGTCATTCCCGACGTCGAGTGTACACACTCCTACTGAGCCTCTTGATGATAAACCCG

 V I S K G C S S Q C E D D S E N Y Y L G

 AAGAAGAACATCACGTGCTGCTACTCTGACCTGTGCAATGTCAACGGGGCCACACCCCTG
 241 -----+-----+-----+-----+-----+ 300
 TTCTTCTGTAGTGCACGACGATGAGACTGGACACGTTACAGTTGCCCGGGTGTGGGAC

 K K N I T C C Y S D L C N V N G A H T L

 AAGCCACCCACCCCTGGGGTGTGCTGACCGTGTCTGACGCTGTTGCTGTGGGGCTCC
 301 -----+-----+-----+-----+-----+ 360
 TTCGGTGGGTGGTGGGACCCGACGACTGGCACGAGACGTCGGACAACGACACCCCGAGG

 K P P T T L G L L T V L C S L L L W G S

 AGCCGTCTGTAGGCTCTGGGAGAGCCTACCATAGCCCGATTGTGAAGGGATGAGCTGCAC
 361 -----+-----+-----+-----+-----+ 420
 TCGGCAGACATCCGAGACCCCTCTCGGATGGTATCGGGCTAACACTTCCCTACTCGACGTG

 S R L *

 TCCACCCACCCCCACACAGG
 421 -----+-----+-----+-----+ 441
 AGGTGGGGTGGGGTGTGTCC

FIG. 3

hSCA-2
hPSCA
mPSCA

1 M K I F L P V L L A A L L G V E R A S S
1 M K A V L L A L L M A G L A L Q P G T A
1 M K T V L F L L L A T Y L A L H P G A A

21 L M C F S C L N Q K S N L Y C L K P T I
21 L L C Y S C K A Q V S N E D C L Q V E N
21 L Q C Y S C T A Q M N N R D C L N V Q N

41 C S D Q D N Y C V T V S A S A G I G N L
41 C T Q L G E Q C W T A R I R A V G L L T
41 C S L D Q H S C F T S R I R A I G L V T

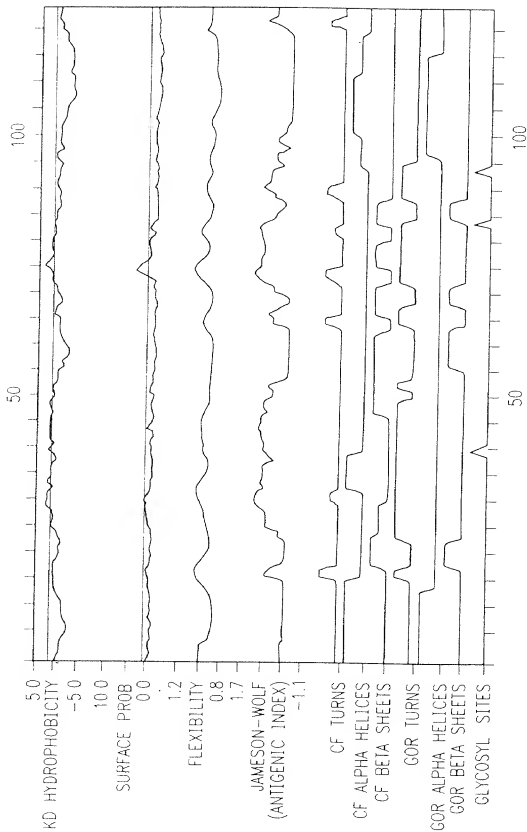
61 V T F G H S L S K T C S P A C P I P E G
61 V - - - - I S K G C S L N C V D D S Q
61 V - - - - I S K G C S S Q C E D D S E

81 V N V G V A S M G I S C C Q S F L C N F
76 D Y Y V G K K - N I T C C D T D L C N A
76 N Y Y L G K K - N I T C C Y S D L C N V

101 S A A D G G L R A S V T L L G A G L L L
95 S G A H A L Q P A A A I L A L L P A L G
95 N G A H T L K P P T T L G L L T V L C S

121 S L L P A L L R F G P
115 L L L W G P G Q L - -
115 L L L W G S S R L L - -

FIG. 4



A schematic diagram of a protein structure. It features a wavy line representing the protein backbone. On the left, a cluster of small circles is labeled "SIGNAL SEQUENCE" with an arrow pointing to it. On the right, a similar cluster is labeled "GPI SIGNAL" with an arrow pointing to it. A legend at the bottom right shows a small circle with a line through it, labeled "= glycosylation SITE".

FIG. 5

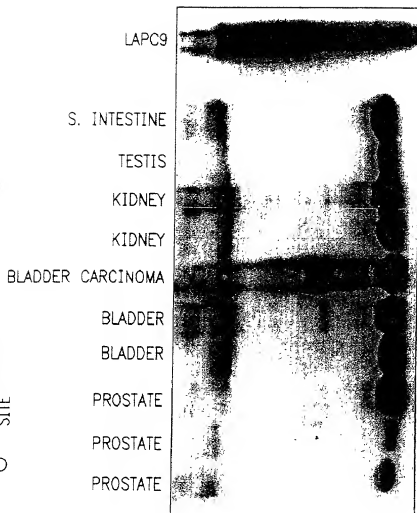


FIG. 6

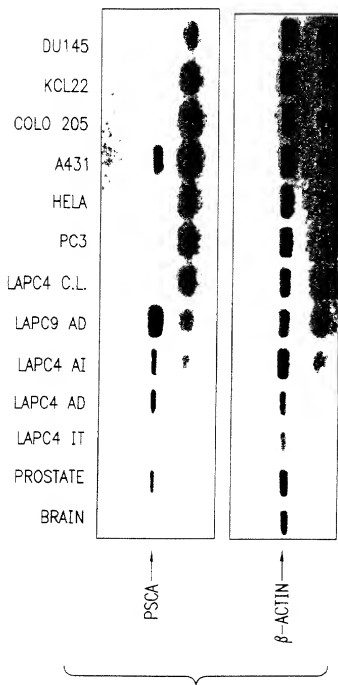
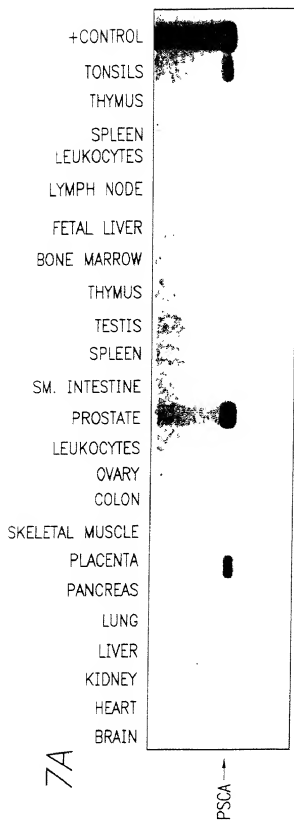


FIG. 8A

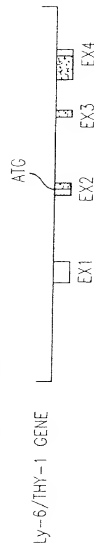


FIG. 8B

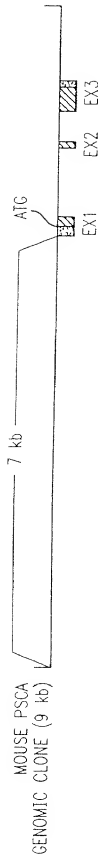


FIG. 8C



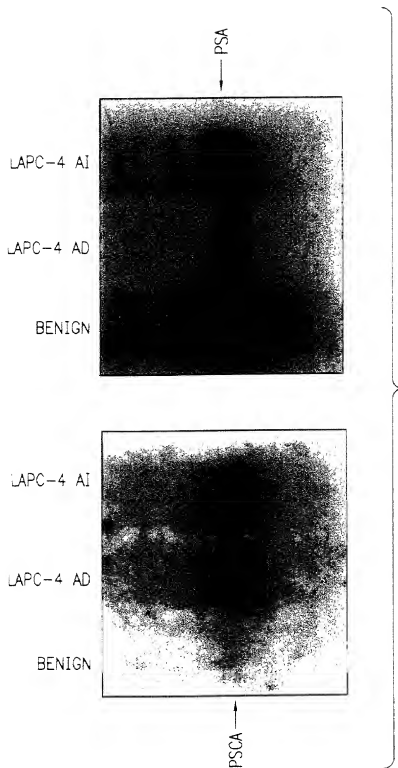


FIG. 9A

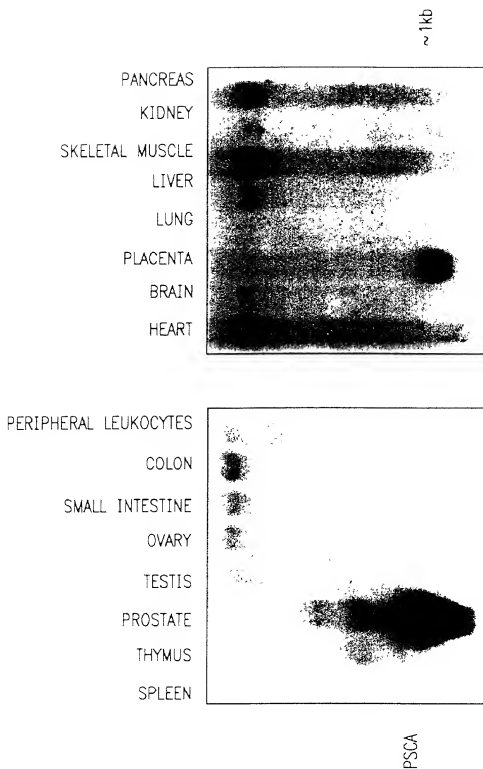


FIG. 9B

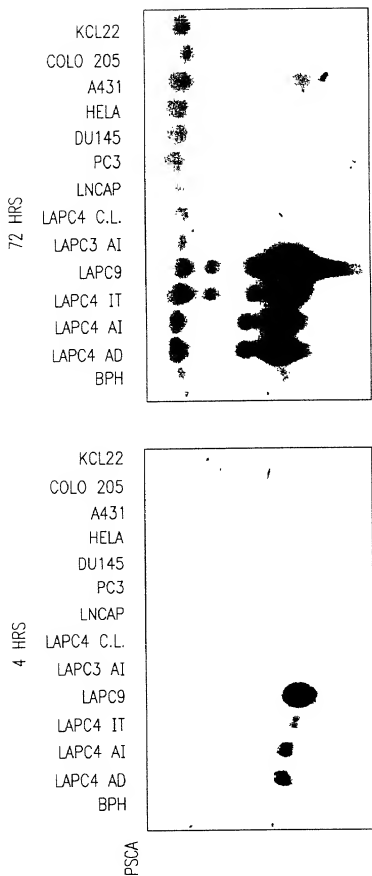


FIG. 10A

	KCL22	
	COLO 205	
	A431	
	HELA	
	DU145	
	PC3	
	LNCAP	
72 HRS	LAPC4 C.L.	
	LAPC3 AI	
	LAPC9	
	LAPC4 IT	
	LAPC4 AI	
	LAPC4 AD	
	BPH	

4 HRS

KCL22
COLO 205
A431
HELA
DU145
PC3
LNCAP
LAPC4 C.L.
LAPC3 AI
LAPC9
LAPC4 IT
LAPC4 AI
LAPC4 AD
BPH

FIG. 10B

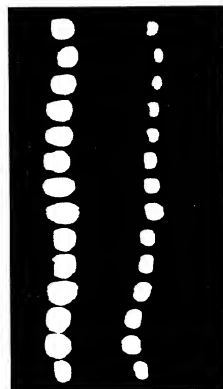
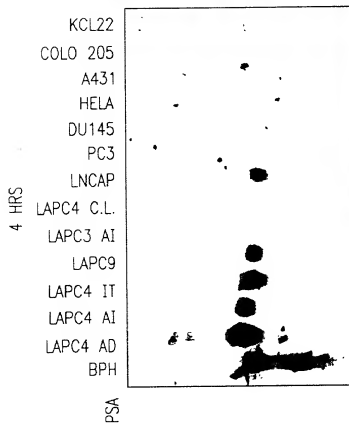
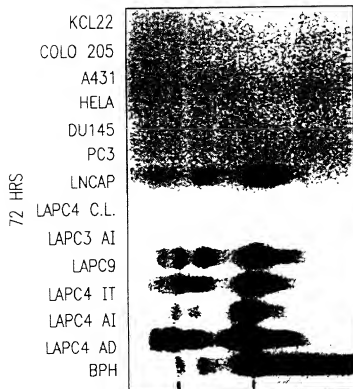


FIG. 10C

FIG. 11A

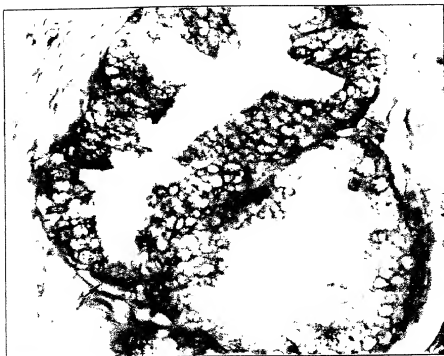


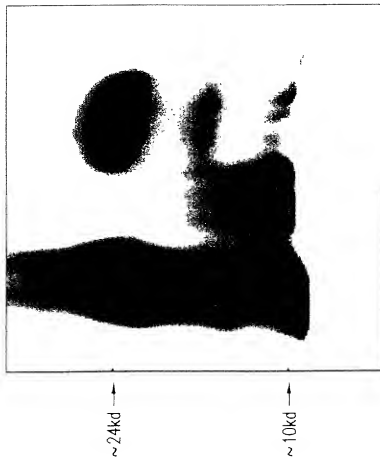
FIG. 11B



FIG. 11C

FIG. 12A

CONTROL
N GLYCOSIDASE F
O GLYCOSIDASE



CELL ASSOCIATED
SECRETED

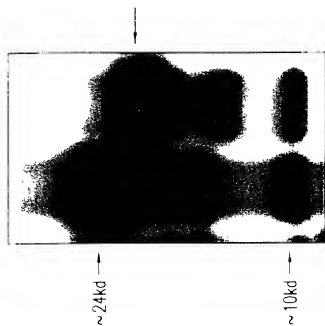


FIG. 12B

FIG. 13

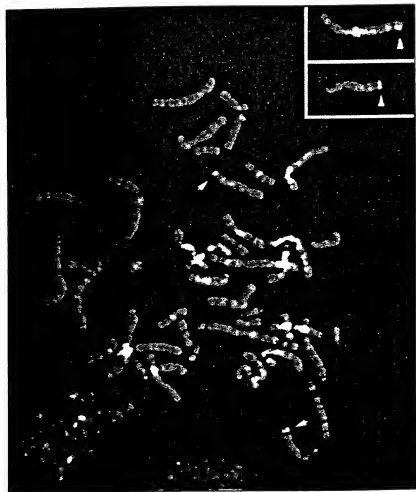


FIG. 14A

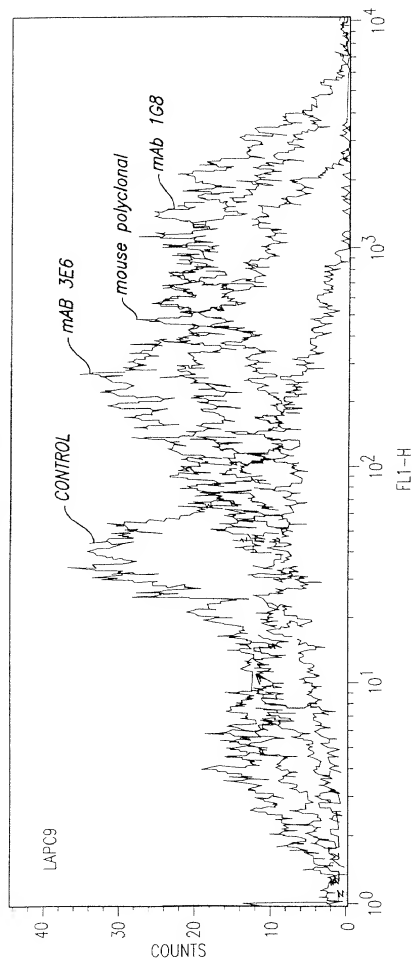


FIG. 14B

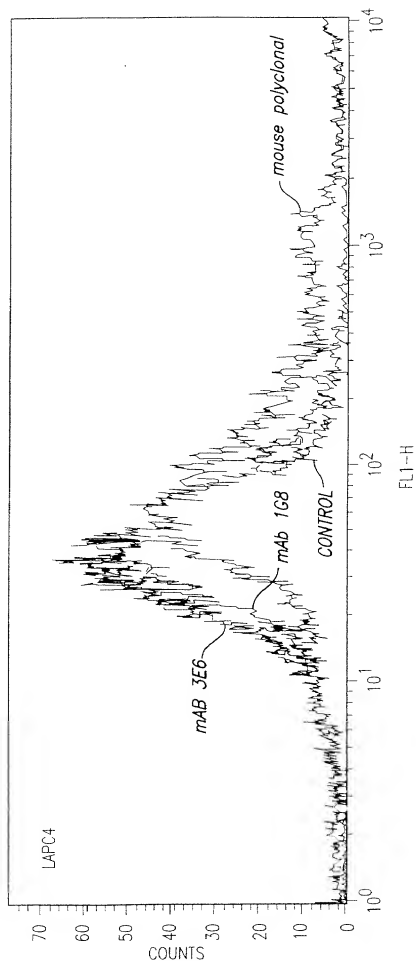
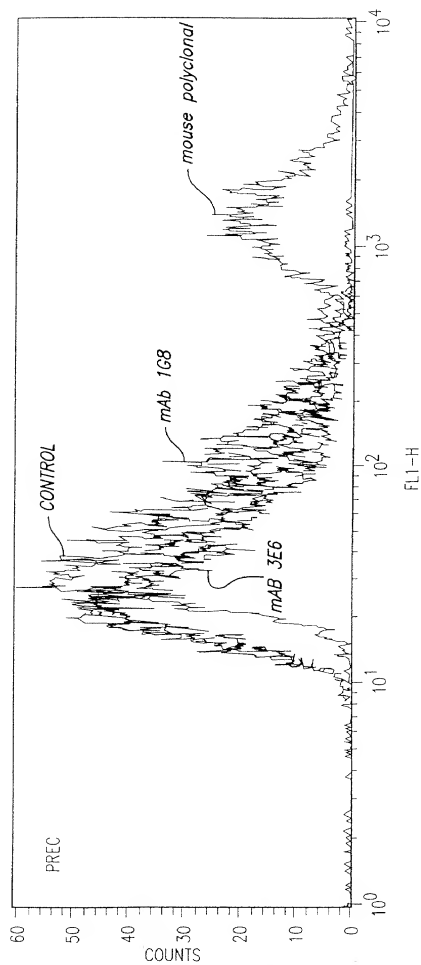


FIG. 14C



mAb	ISOTYPE	EPIOTOPE MAP			
		FL (18-98)	N (2-50)	M (46-109)	C (85-123)
1G8	IgG1 k	2.039	0.007	0.628	0.000
2H9	IgG1 k	1.318	0.863	0.032	0.021
3C5	IgG2a k	2.893	1.965	0.016	0.005
3E6	IgG3 k	0.328	0.024	0.069	0.370
4A10	IgG2a k	2.039	1.315	0.000	0.014
2A2	IgG2a k	1.366	0.733	0.010	0.003
3G3	IgG2a k	2.805	1.731	0.004	0.000

FIG. 15A

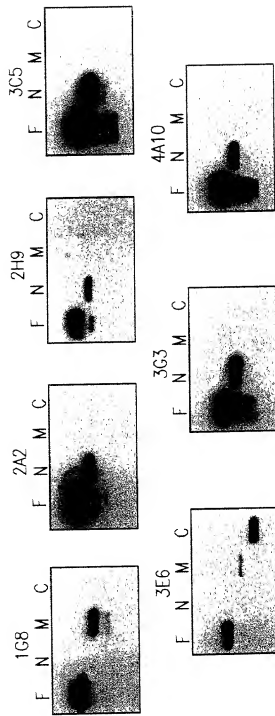


FIG. 15B

PROSTATE STEM CELL ANTIGEN (PSCA) IS A GPI-ANCHORED PROTEIN

1	M	K	I	F	P	V	L	A	A	L	L	G	V	E	R	A	S	hSCA-2
1	M	K	A	V	L	L	A	L	M	A	G	L	A	Q	P	G	T	hPSCA
1	M	K	I	V	F	L	L	A	T	Y	L	A	H	P	C	A	A	mPSCA
21	M	C	F	S	C	L	N	Q	K	S	N	L	Y	C	L	K	P	T
21	L	C	T	S	C	K	A	Q	V	S	N	E	D	C	L	Q	V	E
21	L	Q	C	V	S	C	T	A	Q	M	N	N	R	D	C	L	N	V
41	C	S	D	Q	D	N	Y	C	V	T	V	S	A	S	A	G	I	G
41	C	T	Q	L	G	E	Q	C	W	A	R	T	R	A	V	I	G	L
41	C	S	L	D	Q	H	S	C	F	T	S	R	I	R	A	I	G	L
61	V	T	F	G	H	S	L	S	K	T	C	S	P	A	C	P	I	P
61	V	-	-	-	-	-	-	-	L	S	K	G	C	S	L	N	C	V
61	V	-	-	-	-	-	-	-	L	S	K	G	C	S	S	Q	C	E
81	V	N	V	G	V	A	S	M	G	T	S	C	G	Q	S	F	E	C
76	D	S	V	G	K	K	-	N	I	T	C	G	D	T	D	L	C	N
76	N	-	-	-	-	-	-	-	N	I	T	C	G	Y	S	D	L	C
101	S	A	D	G	G	L	R	A	S	V	T	R	L	G	A	G	T	L
95	S	G	A	H	A	Q	P	A	A	I	L	A	L	P	A	L	G	
95	N	G	A	H	T	L	K	P	P	T	T	L	G	L	T	V	L	C
121	S	E	I	P	A	L	L	R	F	C	P	-	-	-	-	-	-	
115	L	L	W	C	P	G	Q	-	-	-	-	-	-	-	-	-	-	
115	L	L	W	G	S	R	-	-	-	-	-	-	-	-	-	-	-	

FIG. 16A

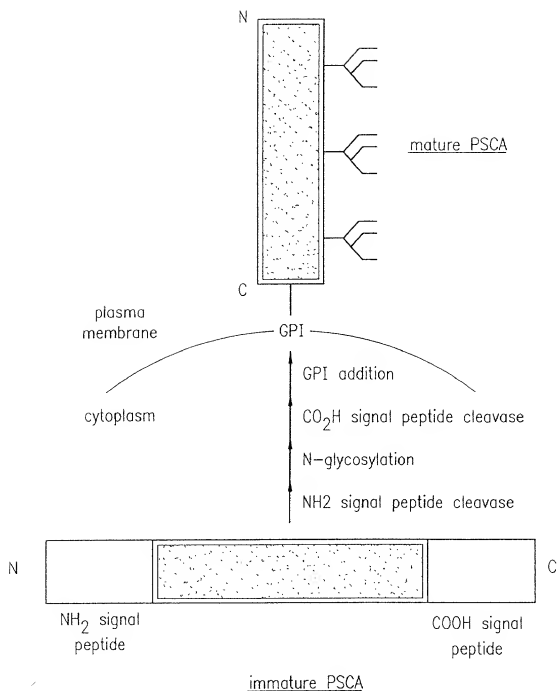


FIG. 16B

FIG. 17

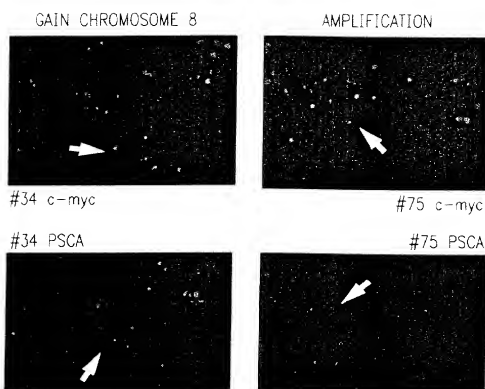


FIG. 18

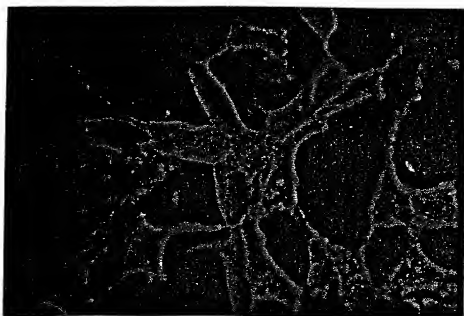




FIG. 19

FIG. 20

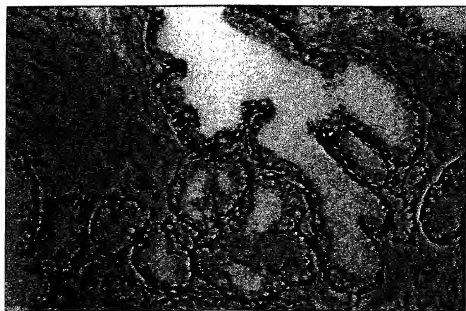


FIG. 21

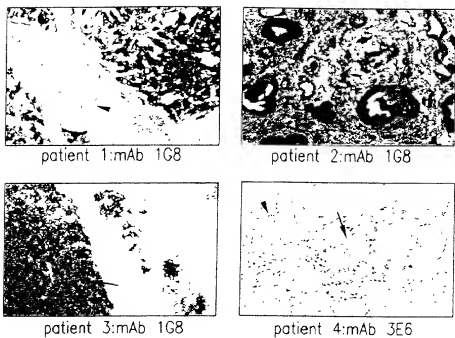
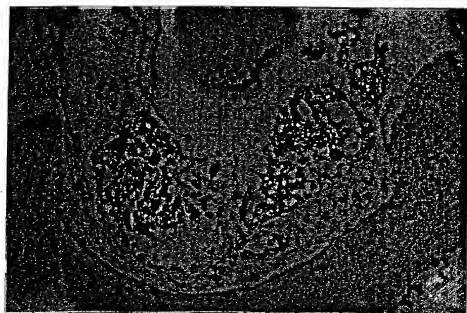


FIG. 22



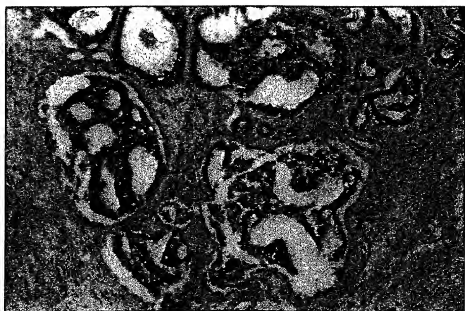


FIG. 23

FIG. 24



FIG. 25

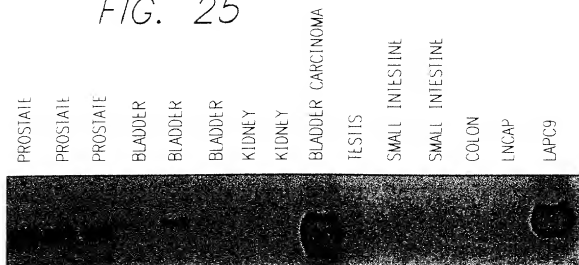


FIG. 26

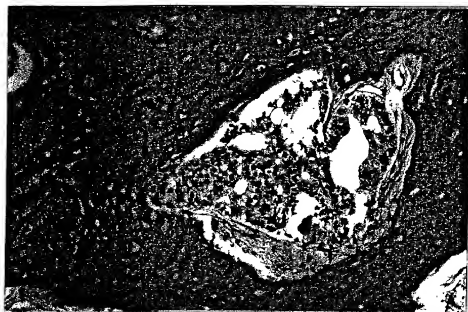
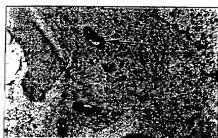


FIG. 27



Patient 5: H and E
and mAb 1G8



Patient 4: H and E
and mAb 3E6

FIG. 28

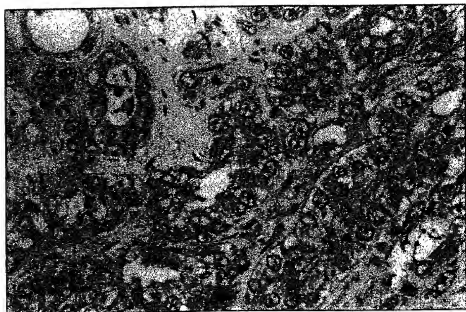
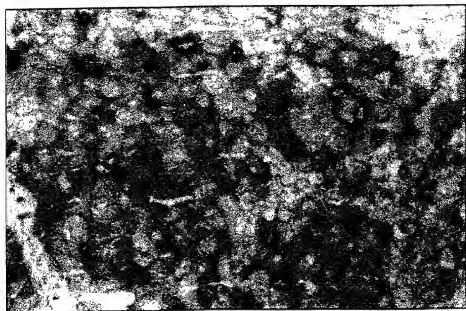


FIG. 29

FIG. 30



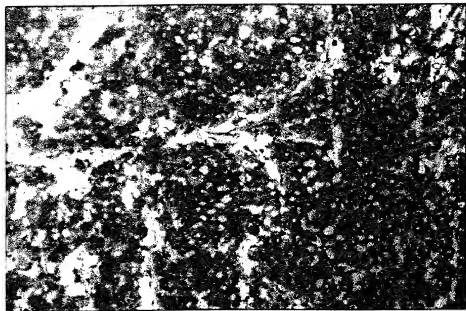


FIG. 31

FIG. 32

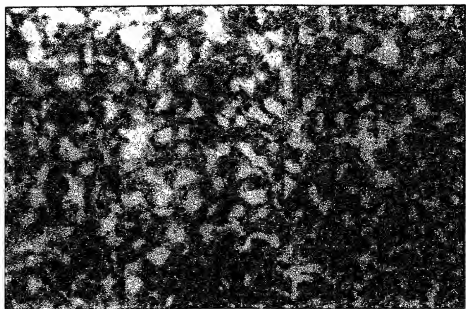


FIG. 33

SECONDARY ANTIBODY

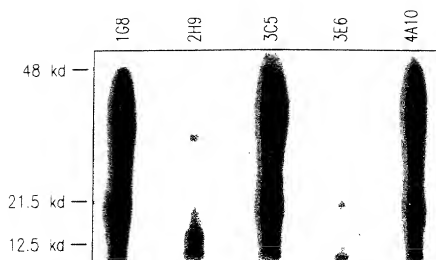
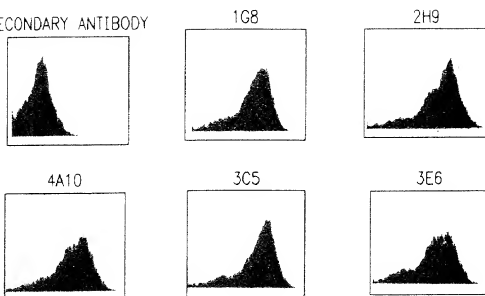


FIG. 34

FIG. 35

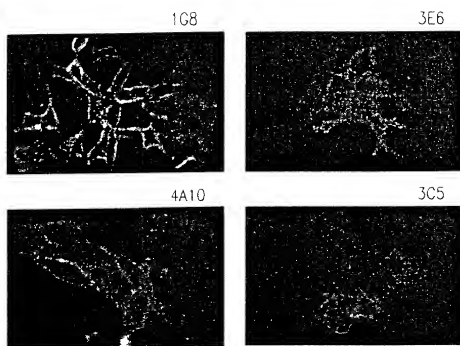


FIG. 36

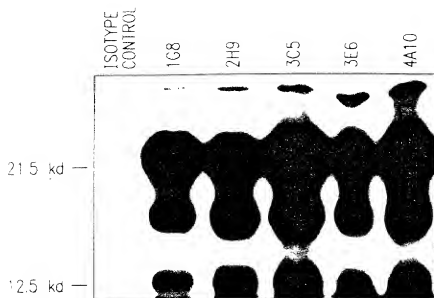


FIG. 37

NORMAL: ISOTYPE CONTROL



NORMAL: PSCA mAb 3E6



NORMAL: PSCA mAb 1G8



ATROPHY: PSCA mAb 2H9

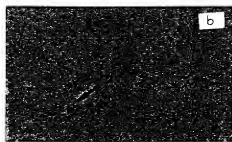


FIG. 38

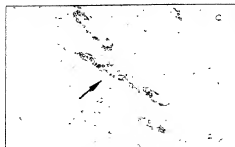
FIG. 39A



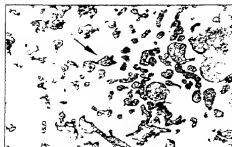
BLADDER: 1G8



COLON: 1G8



KIDNEY: 3E6



PLACENTA: 3E6

PROSTATE

PROSTATE

PROSTATE

KIDNEY

KIDNEY

KIDNEY

BLADDER

BLADDER

BLADDER

LAPC 9



PSCA



ACTIN

FIG. 39B

FIG. 40A

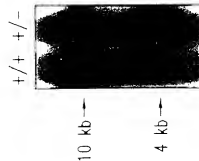
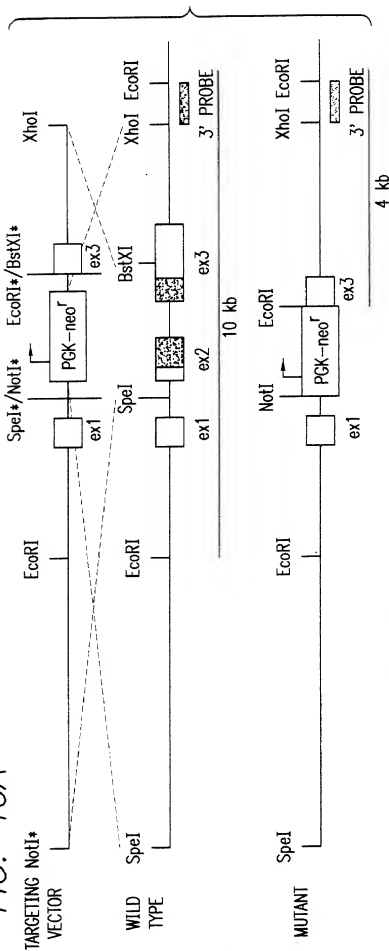


FIG. 40B

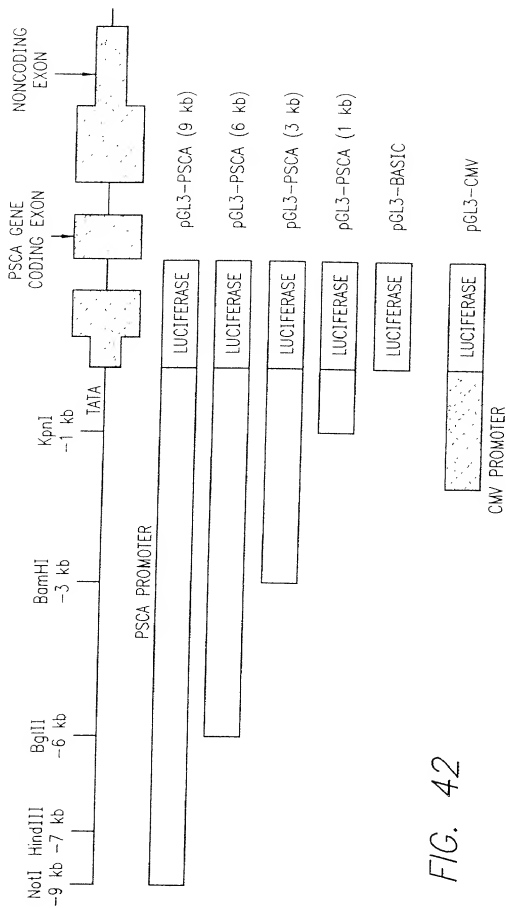


FIG. 42

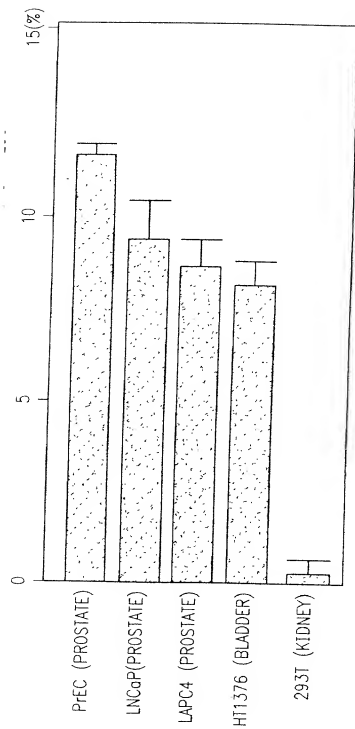


FIG. 43

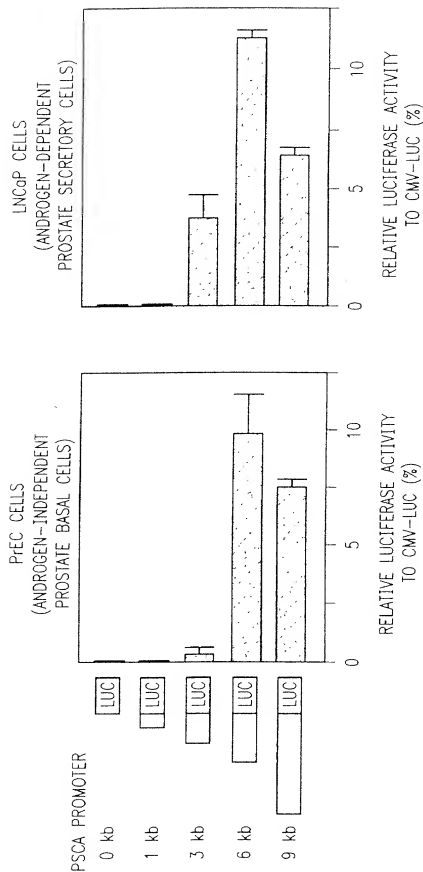
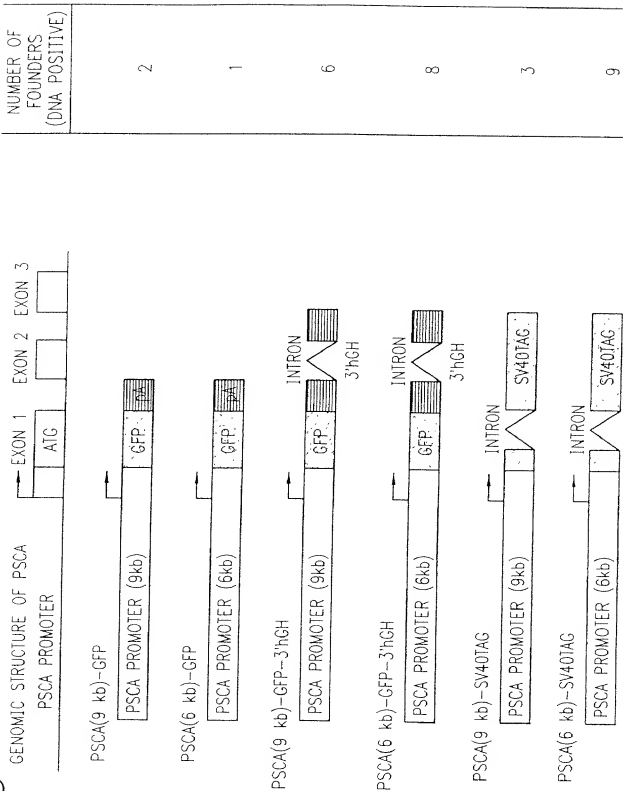


FIG. 44

FIG. 45



NEGATIVE TISSUES

STOMACH
SMALL INTESTINE
COLON
SEMINAL VESICLE
URETHRA
TESTIS
LIVER
KIDNEY
LUNG
BRAIN
HEART
SKELETAL MUSCLE
OVARY
UTERUS

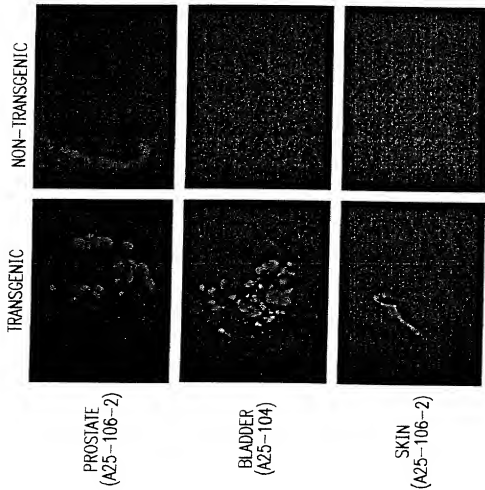


FIG. 46

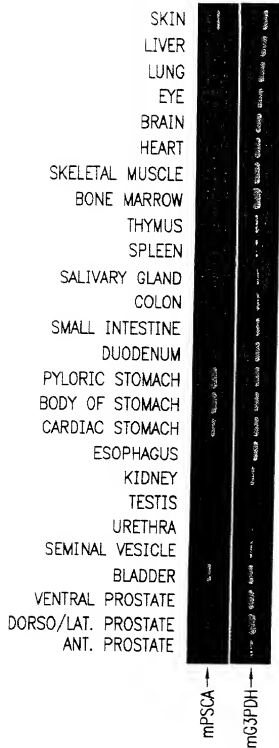
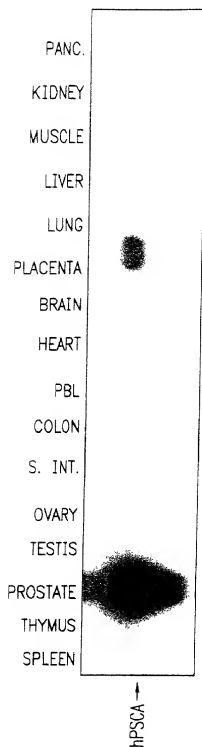


FIG. 47

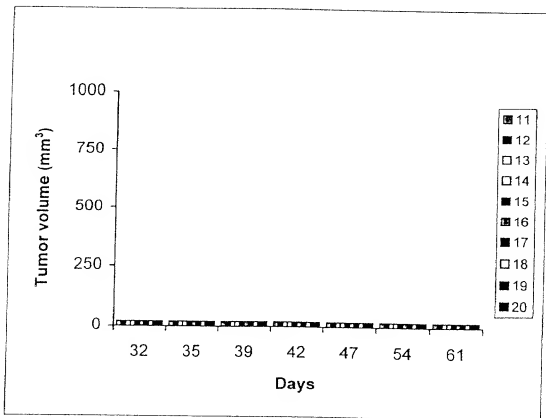
[illegible]

FIG. 49

A

Epitope recognized (OD 450 nm)

mAb	Isotype	F (18-98)	N (2-50)	M (46-109)	C (85-123)
1G8	IgG1 k	1.485	0.004	1.273	0.003
2A2	IgG2a k	0.973	0.631	0.023	0.010
2H9	IgG1 k	1.069	1.026	0.002	0.001
3C5	IgG2a k	1.916	1.709	0.006	0.002
3E6	IgG3 k	1.609	0.036	1.133	2.118
3G3	IgG2a k	2.805	1.731	0.004	0.000
4A10	IgG2a k	1.053	0.493	0.000	0.001

B

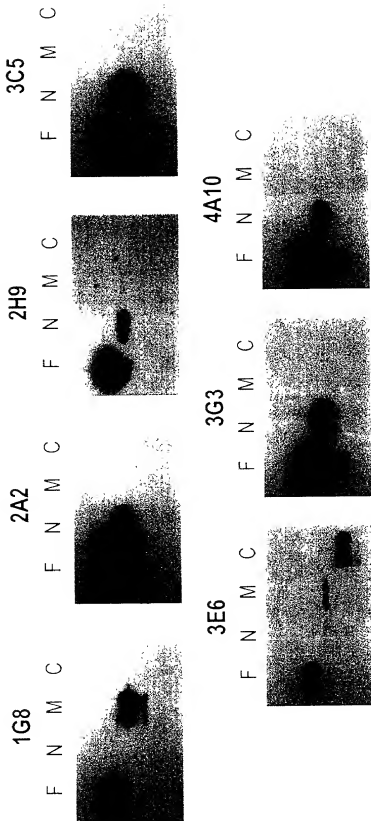
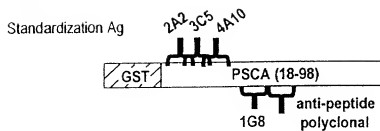
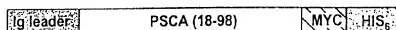


FIG. 50

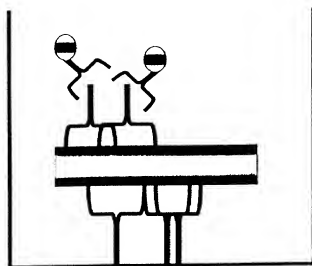
A



Engineered mammalian secreted form



B



Anti-IgG2a HRP

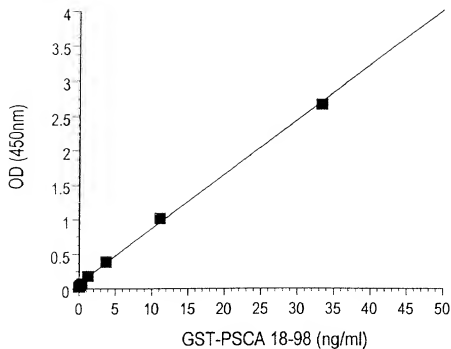
Anti-PSCA mAbs 3C5+4A10+2A2 (IgG2a)

PSCA

Affinity purified anti-peptide polyclonal
+ mAb 1G8 (IgG1)

FIG. 51

A



B

Sample	OD+range (n=2)	ng/ml
vector	0.005+0.001	ND
vector+hu serum	0.004+0.001	ND
secPSCA	2.695+0.031	32.92
secPSCA+hu serum	2.187+0.029	26.55

FIG. 52

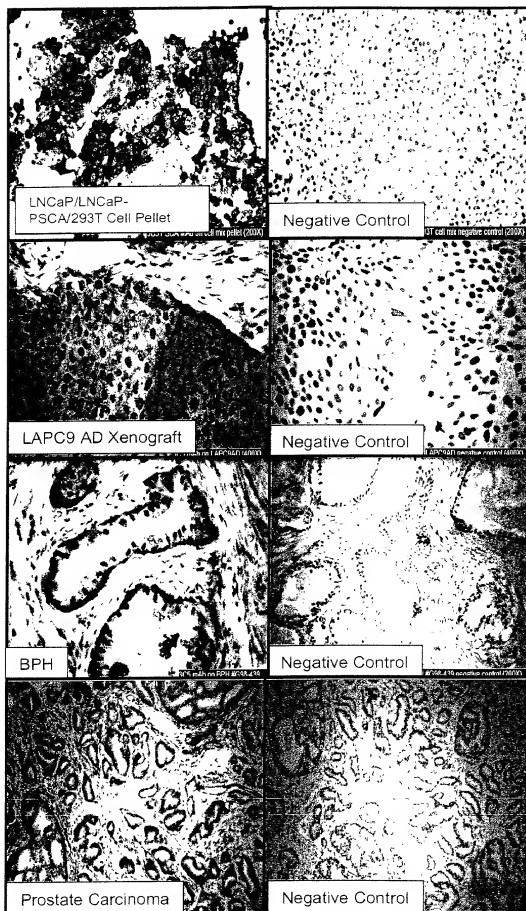


FIG. 53

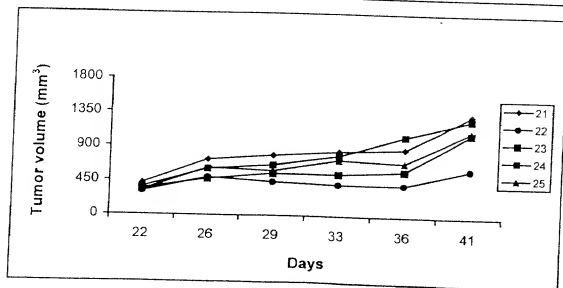
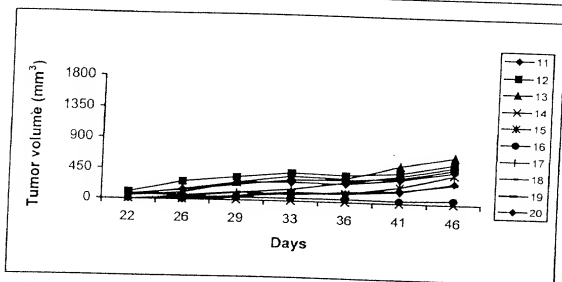
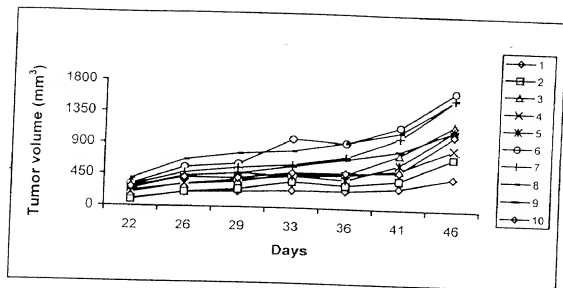


FIG. 54

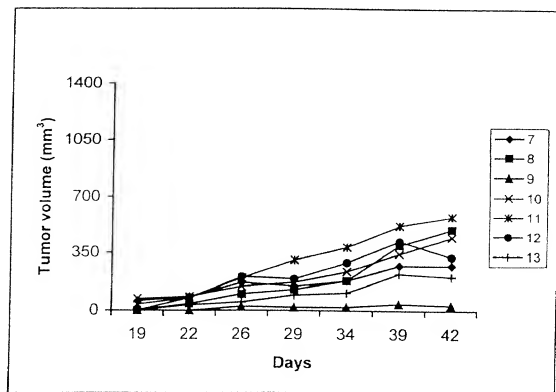
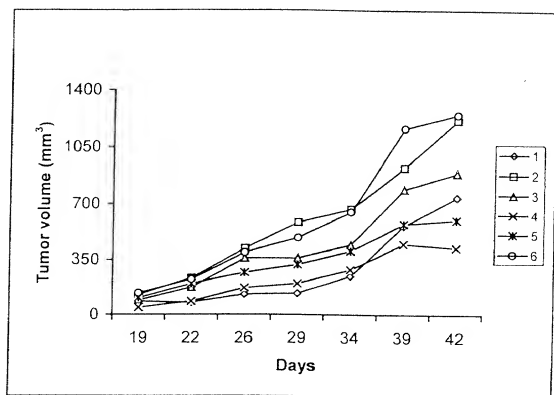


FIG. 55

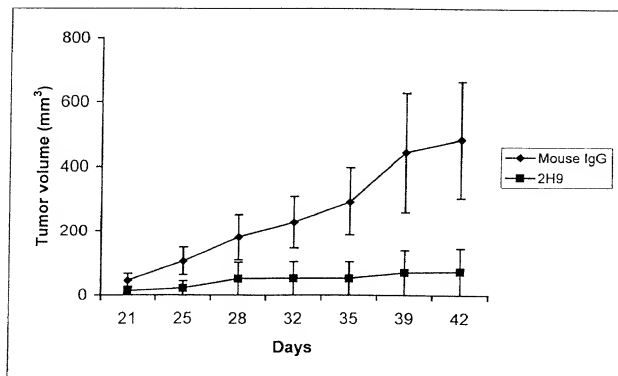
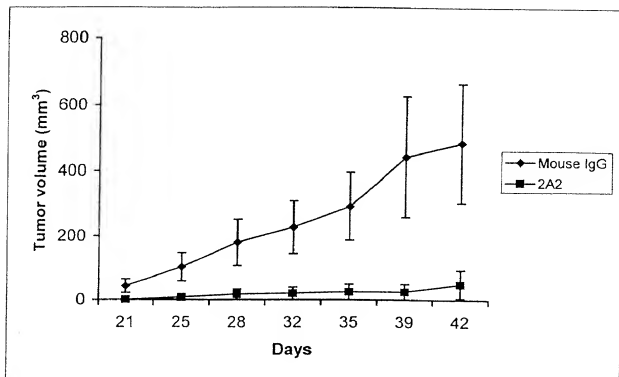


FIG. 56

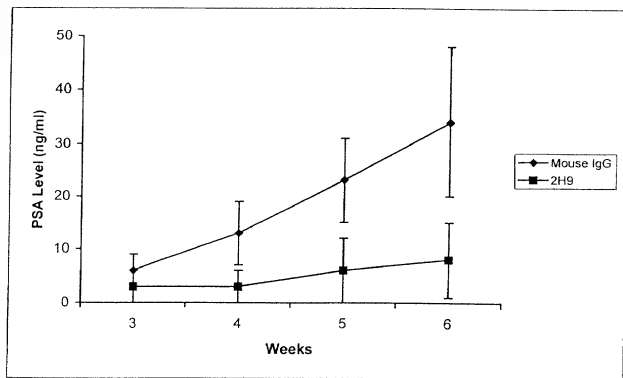
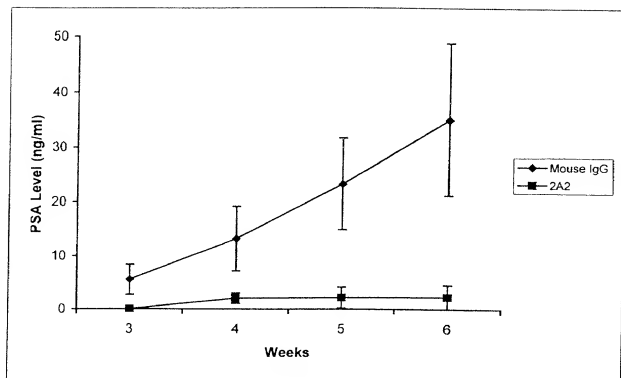


FIG. 57

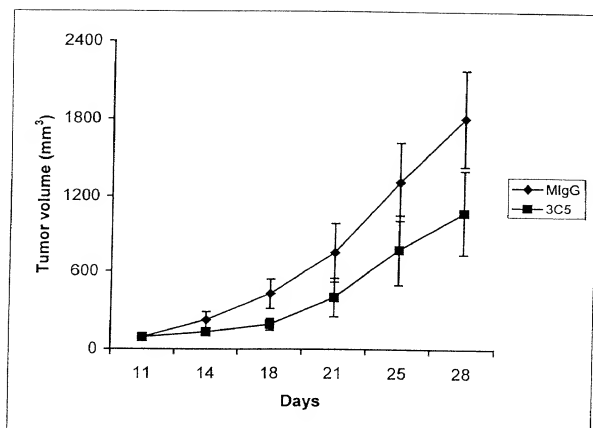


FIG. 58

TCTCTCTTCTGAGTGGCACTGGTTATAGAGTCAATTCAGAGGTTTCAGCTGCAGCAGTCT 60
 C F F L M A V V I G V N S E V Q L Q Q S 20

 GGGGCGAGAACTTGTGAGGTCAGGGGCTCAGTCAAGTTGTCTCTGCACAGCTTCTGGCTTC 120
 G A E L V R S G A S V K L S C T A S G F 40

 AACTTTAAAGACTACTATATACACTGGGTGAATCAGAGGCTGACCCAGGGCTGGAGTGG 180
 N I K D Y Y I H W V N Q R P D Q G L E W 60

 ATTGGATGGATTGATCTCTGAGAATGGTGACACTGAATTGTCTCCGAAGTTCAGGGCAAG 240
 I G W I D P E N G D T E F V P K F Q G K 80

 GCCACTATGACTGCAGACATTCTCTCCACACAGCCTACCTGCACCTCAGCAGCCTGACA 300
 A T M T A D I F S N T A Y L H L S S L T 100

 TCTGAAGACACTGCCGTCTATTACTGTAAACGGGGGTTTCTGGGGCCAAAGGACTCTG 360
 S E D T A V Y Y C K T G G F W G Q G T L 120

 GTCATGTCTCTGCAGCCAAACGACACCCCATCTGTATCCACTG
 V T V S A A K T T P P S V Y P L

FIG. 59

TTGGTAGCAACAGCCTCAGATGCCACTCCAGGTCCAACTGCAGCAACCTGGGCTGAA 60
L V A T A S D V H S Q V Q L Q Q P G S E 20

CTGGTAGGCGCTGGAACTTCAGTGAAGCTCTCTGCAAGGCTTCCTGGCTATACATCTCC 120
L V R P G T S V K L S C K A S G Y T F S 40
CDR1

AGTACTGGATGCACCTGGGTGAGCAGAGCGCTGGACAGGCCTTGAGTGGATTGGAAT 180
S Y W M H W V K Q R P G Q G L E W I G N 60

ATTGACCTGGTAGTGTTACACTAATACGCTGAGAACCCTCAAGACCAAGGCCACATG 240
I D P G S G Y T N Y A E N L K T K A T L 80
CDR2

ACTGTAGACACATCCTCCAGCACAGCCTACATGCAGCTCAGCAGCCTGCATCTGAGGAC 300
T V D T S S S T A Y M Q L S S L T S E D 100

TCTGCAGTCTATTACTGTACAAGCGCATCTACTATGATTACGACGGGATTGCTTACTGG 360
S A V Y Y C T S R S T M I T T G F A Y W 120
CDR3

GGCCAGGGACTCTGGTCACGTCTCTGCGAGCTACACACAGCCCACTCTGTATCCA 420
G Q G T L V T V S A A T T A P S V Y P 160

CTGGCC
L A

FIG. 60

ANTGACTTCGGGTGAGCTGGGTTTTATTATTGTCTTTTAAAGGGTCCGAGTGAA 60
N D F G L S W V F I I V L L K G V R S E 20

GTGAGGCTTGAGGAGTCTGGAGGAGCTGGTGCAACTGGAGGATCCATGAACTCTCC 120
V R L E E S G G G W V Q P G G S M K L S 40

TGTGTAGCCTCTGGATTCTTACTTTTACGTAAATTACTGGATGACTTGGTCCGCCAGTCTCCA 180
C V A S G F T F S N Y W M T W V R Q S P 60
CDR1

GAGAAAGGGCTTGAGTGGGTTGCTGAAATTCGATTGAGATCTGAAATTTATGCAACACAT 240
E K G L E W V A E I R L R S E N Y A T H 80
CDR2

TATCGGAGTCTGTGAAGGGAATTCACCATCTCAAGAGATGATCCAGAAGTCGTCTC 300
Y A E S V K G K F T I S R D D S R S R L 100

TACCTGCAAAATGAACAACCTTAAGACCTGAAGACAGTGGAAATTTATTACTGTACAGATGGT 360
Y L Q M N N L R P E D S G I Y Y C T D G 120

CTGGACGACCTTACTGGGGCCAGGAGCTGTGTCACGTCTCTGAGGCCAAACGACA 420
L G R P N W G Q G T L V T V S A A K T T 140
CDR3

CCCCATCTGTCTATCCACTGGCCCTTGTGTA
P P S V Y P L A P C V

FIG. 61

CDR1 Comparisons

1G8	1gG _{1k}	Middle	G	F	N	I	K	D	Y	Y	I	H
2H9	1gG _{1k}	N-Term.	G	F	T	F	S	N	Y	W	M	T
4A10	1gG _{2ak}	N-Term.	G	Y	T	F	S	S	Y	W	M	H

CDR2 Comparisons

1G8	1gG _{1k}	W	I	D	P	E	N	G	D	T	E	F	V	P	K	F	Q	G		
2H9	1gG _{1k}	E	I	R	L	R	S	E	N	Y	A	T	H	Y	A	E	S	V	K	G
4A10	1gG _{2ak}	N	I	D	P	G	S	G	Y	T	N			Y	A	E	N	L	K	T

CDR3 Comparisons

1G8	1gG _{1k}	G	G	F														
2H9	1gG _{1k}	L	G	R	P	N												
4A10	1gG _{2ak}	R	S	T	M	I	T	T	G	F	A	Y						

FIG. 62

A



B



C



D

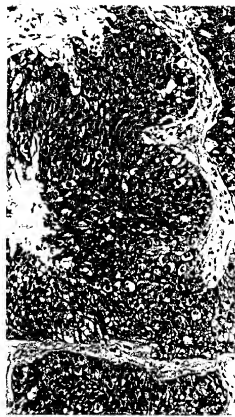


FIG. 63

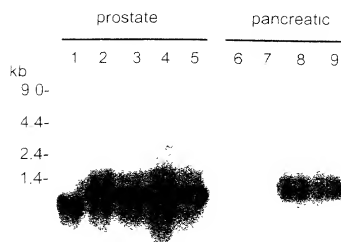


FIG. 64

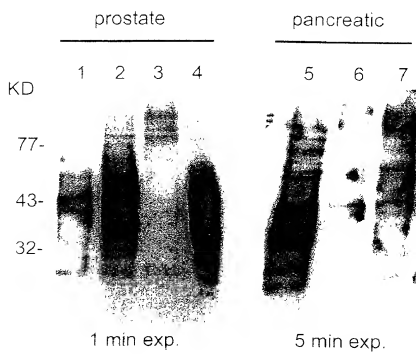


FIG. 65

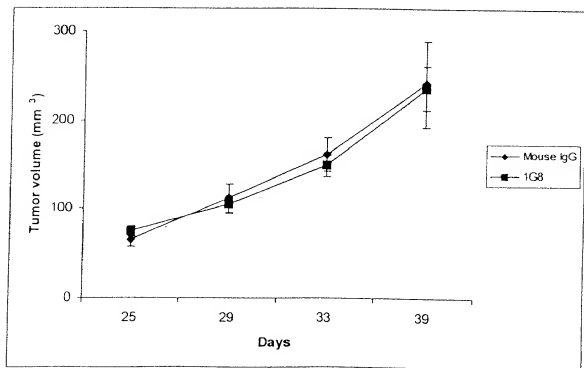
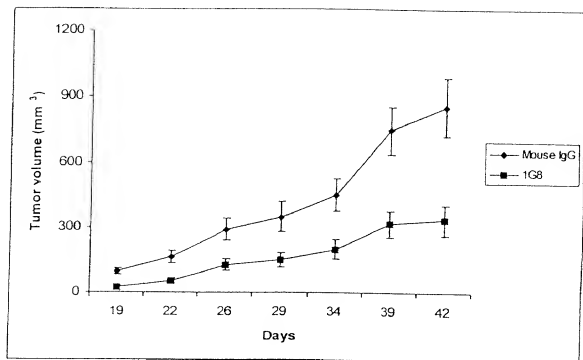
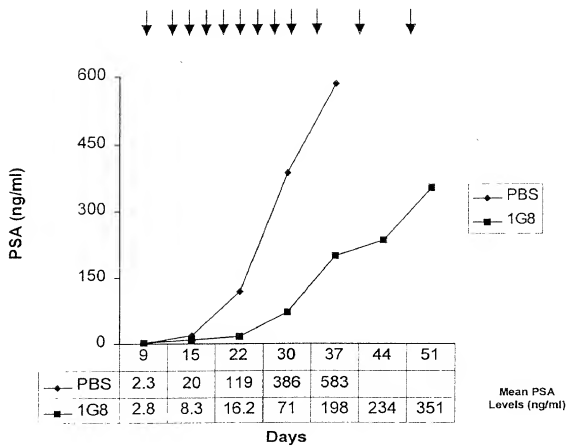


FIG. 66

A)



B)

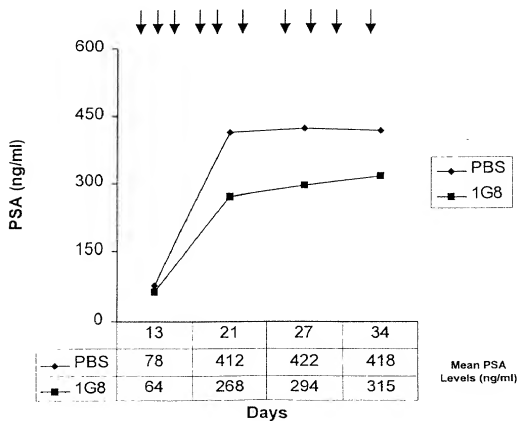
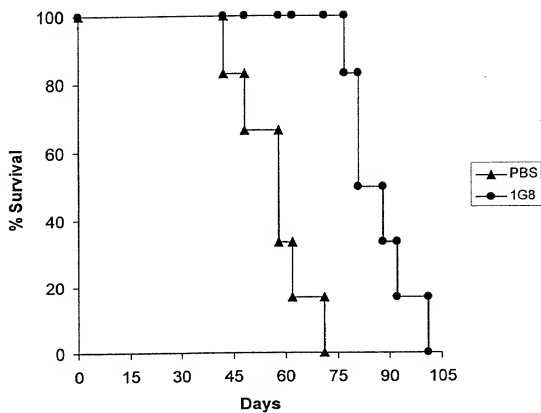


FIG. 67

A)



B)

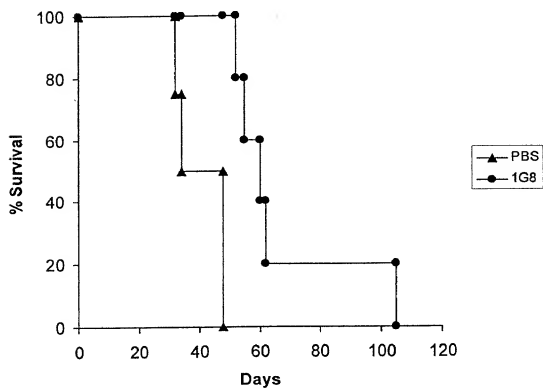
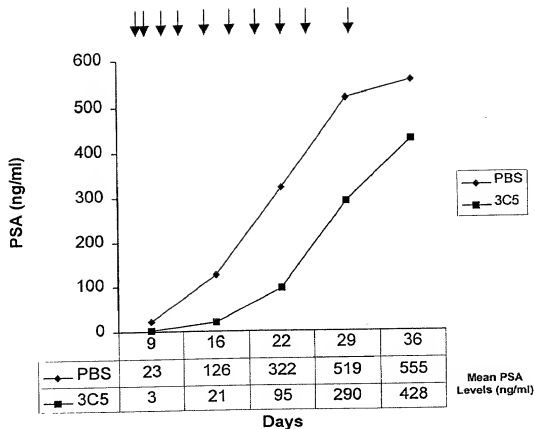


FIG. 68

A)



B)

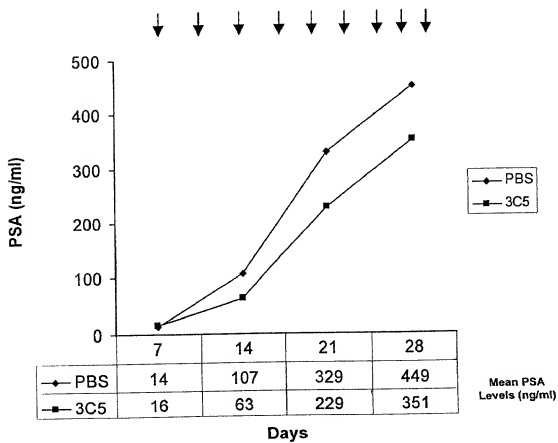
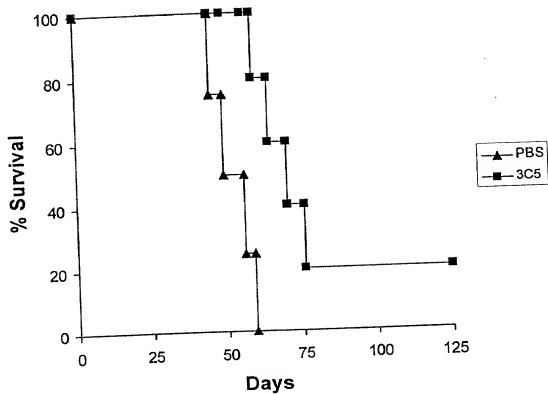


FIG. 69

A)



B)

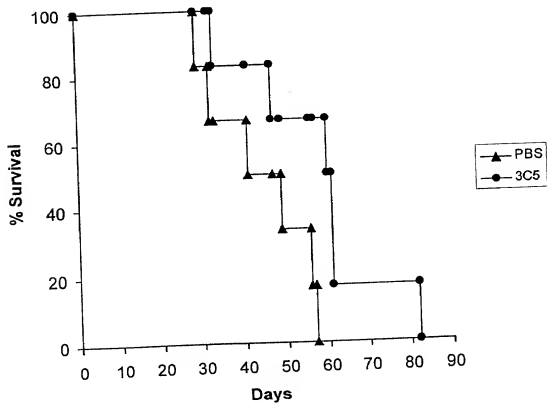


FIG. 70

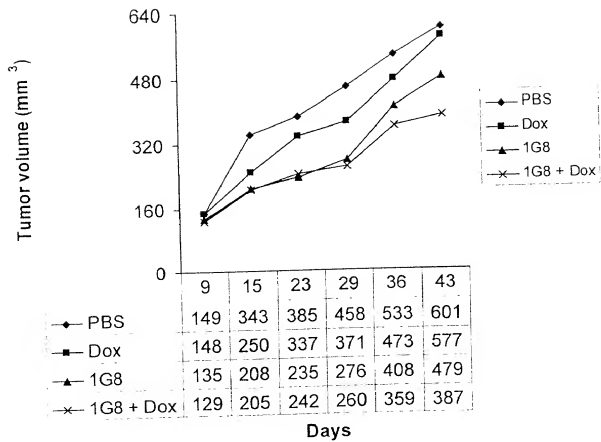
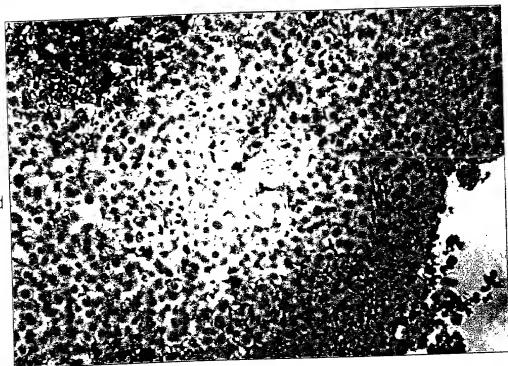
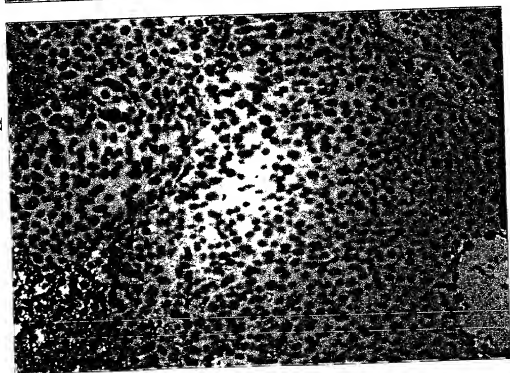


FIG. 71

3C5 Treated



mIgG Treated



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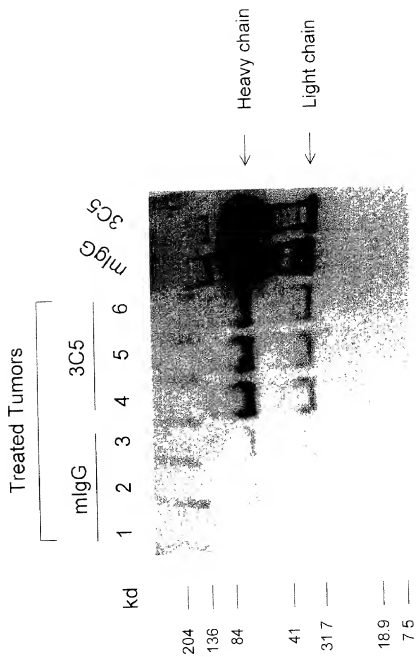


FIG. 72

7G8
mlgG

	mlgG Treated	1G8 Treated
1	0.00	0.00
2	0.00	0.00
3	0.00	0.00
4	0.00	0.00
5	0.00	0.00
6	0.00	0.00
7	0.00	0.00
8	0.00	0.00
9	0.00	0.00
10	0.00	0.00
11	0.00	0.00
12	0.00	0.00
13	0.00	0.00
14	0.00	0.00
15	0.00	0.00
16	0.00	0.00
17	0.00	0.00
18	0.00	0.00
19	0.00	0.00
20	0.00	0.00
21	0.00	0.00
22	0.00	0.00
23	0.00	0.00
24	0.00	0.00
25	0.00	0.00
26	0.00	0.00
27	0.00	0.00
28	0.00	0.00
29	0.00	0.00
30	0.00	0.00
31	0.00	0.00
32	0.00	0.00
33	0.00	0.00
34	0.00	0.00
35	0.00	0.00
36	0.00	0.00
37	0.00	0.00
38	0.00	0.00
39	0.00	0.00
40	0.00	0.00
41	0.00	0.00
42	0.00	0.00
43	0.00	0.00
44	0.00	0.00
45	0.00	0.00
46	0.00	0.00
47	0.00	0.00
48	0.00	0.00
49	0.00	0.00
50	0.00	0.00
51	0.00	0.00
52	0.00	0.00
53	0.00	0.00
54	0.00	0.00
55	0.00	0.00
56	0.00	0.00
57	0.00	0.00
58	0.00	0.00
59	0.00	0.00
60	0.00	0.00
61	0.00	0.00
62	0.00	0.00
63	0.00	0.00
64	0.00	0.00
65	0.00	0.00
66	0.00	0.00
67	0.00	0.00
68	0.00	0.00
69	0.00	0.00
70	0.00	0.00
71	0.00	0.00
72	0.00	0.00
73	0.00	0.00
74	0.00	0.00
75	0.00	0.00
76	0.00	0.00
77	0.00	0.00
78	0.00	0.00
79	0.00	0.00
80	0.00	0.00
81	0.00	0.00
82	0.00	0.00
83	0.00	0.00
84	0.00	0.00
85	0.00	0.00
86	0.00	0.00
87	0.00	0.00
88	0.00	0.00
89	0.00	0.00
90	0.00	0.00
91	0.00	0.00
92	0.00	0.00
93	0.00	0.00
94	0.00	0.00
95	0.00	0.00
96	0.00	0.00
97	0.00	0.00
98	0.00	0.00
99	0.00	0.00
100	0.00	0.00

6

204	—
136	—
84	—
41	—
317	—
189	—
75	—

← Heavy chain

← Light chain

FIG. 73